endo sequence from Collagen XVIII.

Sequence Range: 1-555

Nucleotide 1 = Start for Endostatin and fragments EM1 and EM2.

EM1 fragment ends at nucleotide 525, EM2 fragment ends at nucleotide 501.

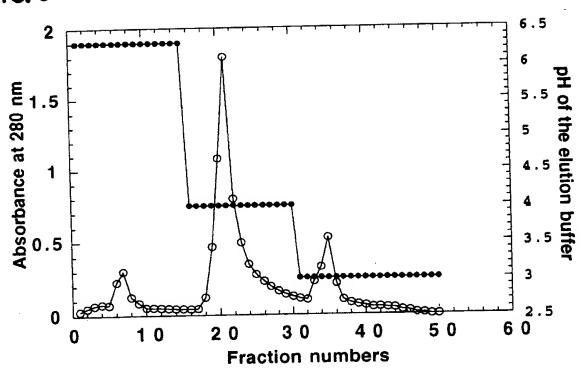
CAT ACT CAT CAG GAC TTT CAG CCA GTG CTC CAC CTG GTG GCA CTG AAC GTA TGA GTA GTC CTG AAA GTC GGT CAC GAG GTG GAC CAC CGT GAC TTG ACC CCC CTG TCT GGA GGC ATG CGT GGT ATC CGT GGA GCA GAT TTC CAG TGG GGG GAC AGA CCT CCG TAC GCA CCA TAG GCA CCT CGT CTA AAG GTC TGC TTC CAG CAA GCC CGA GCC GTG GGG CTG TCG GGC ACC TTC CGG GCT ACG AAG GTC GTT CGG GCT CGG CAC CCC GAC AGC CCG TGG AAG GCC CGA TTC CTG TCC TCT AGG CTG CAG GAT CTC TAT AGC ATC GTG CGC CGT GCT AAG GAC AGG AGA TCC GAC GTC CTA GAG ATA TCG TAG CAC GCG GCA CGA GAC CGG GGG TCT GTG CCC ATC GTC AAC CTG AAG GAC GAG GTG CTA TCT CTG GCC CCC AGA CAC GGG TAG CAG TTG GAC TTC CTG CTC CAC GAT AGA CCC AGC TGG GAC TCC CTG TTT TCT GGC TCC CAG GGT CAA CTG CAA CCC GGG TCG ACC CTG AGG GAC AAA AGA CCG AGG GTC CCA GTT GAC GTT GGG GGG GCC CGC ATC TTT TCT TTT GAC GGC AGA GAT GTC CTG AGA CAC CCA CCC CGG GCG TAG AAA AGA AAA CTG CCG TCT CTA CAG GAC TCT GTG GGT GCC TGG CCG CAG AAG AGC GTA TGG CAC GGC TCG GAC CCC AGT GGG CGG CGG ACC GGC GTC TTC TCG CAT ACC GTG CCG AGC CTG GGG TCA CCC GCC AGG CTG ATG GAG AGT TAC TGT GAG ACA TGG CGA ACT GAA ACT ACT GGG TCC GAC TAC CTC TCA ATG ACA CTC TGT ACC GCT TGA CTT TGA TGA CCC GCT ACA GGT CAG GCC TCC TCC CTG CTG TCA GGC AGG CTC CTG GAA CAG CGA TGT CCA GTC CGG AGG AGG GAC GAC AGT CCG TCC GAG GAC CTT GTC AAA GCT GCG AGC TGC CAC AAC AGC TAC ATC GTC CTG TGC ATT GAG AAT TTT CGA CGC TCG ACG GTG TTG TCG ATG TAG CAG GAC ACG TAA CTC TTA AGC TTC ATG ACC TCT TTC TCC AAA TAG TCG AAG TAC TGG AGA AAG AGG TTT ATC

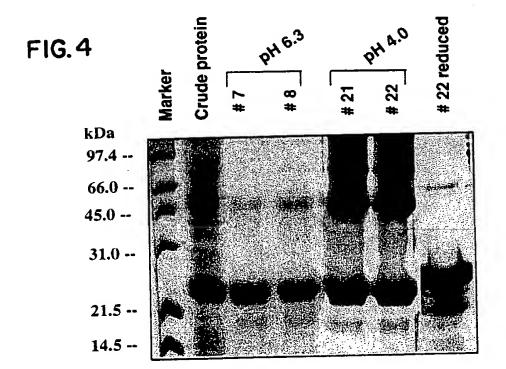
Sequence Range: 1 to 184

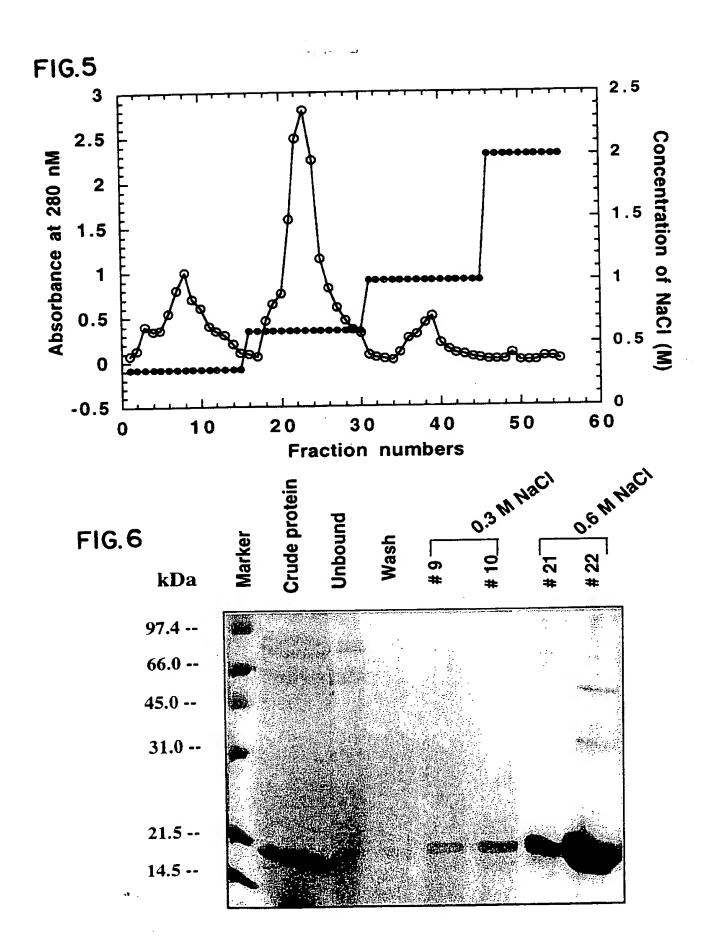
HTH QDF QPV LHL VAL NTP LSG GMR GIR GAD FQC FQQ ARA VGL SGT FRA FLS SRL QDL YSI VRR ADR GSV PIV NLK DEV LSP SWD SLF SGS QGQ LQP GAR IFS FDG RDV LRH PAW PQK SVW HGS DPS GRR LME SYC ETW RTE TTG ATG QAS SLL SGR LLE QKA ASC HNS YIV LCI ENS FMT SFS K

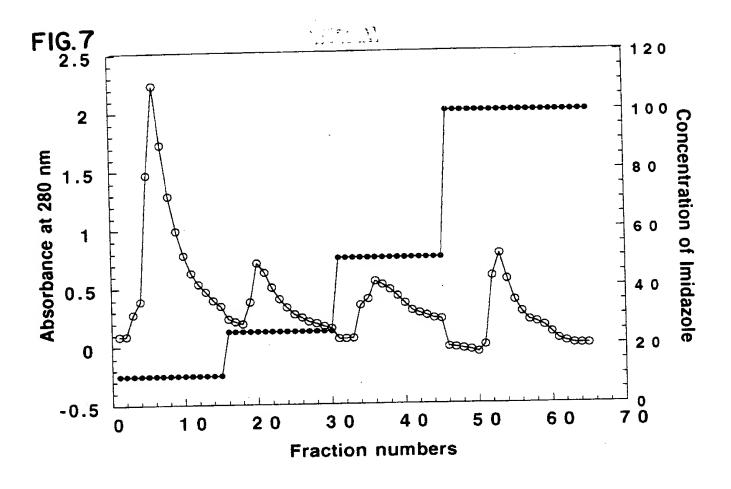
Fig. 2

FIG. 3









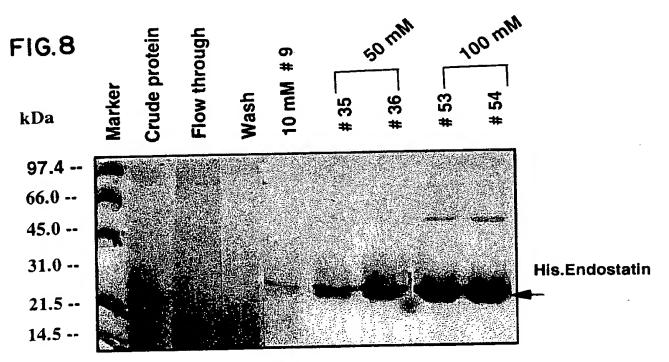
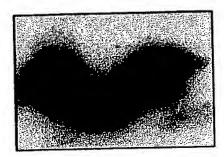
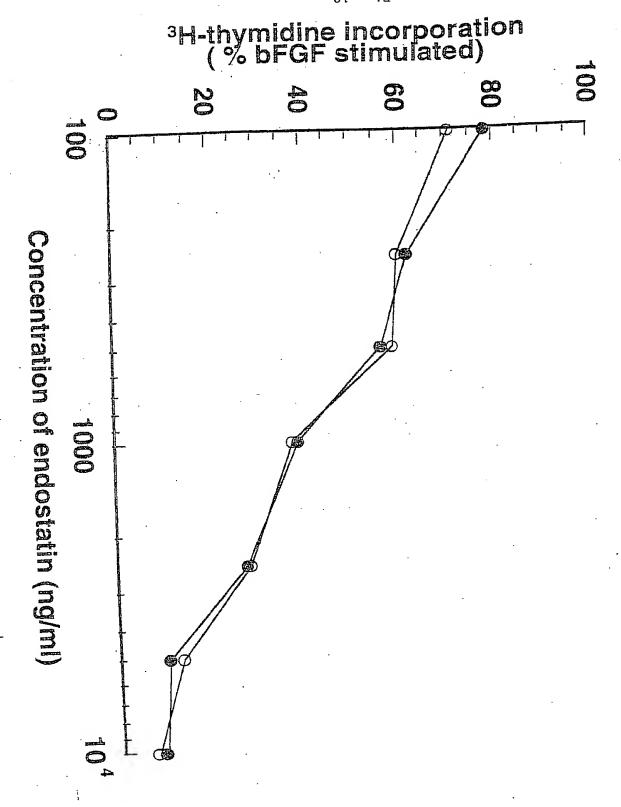


FIG. 9

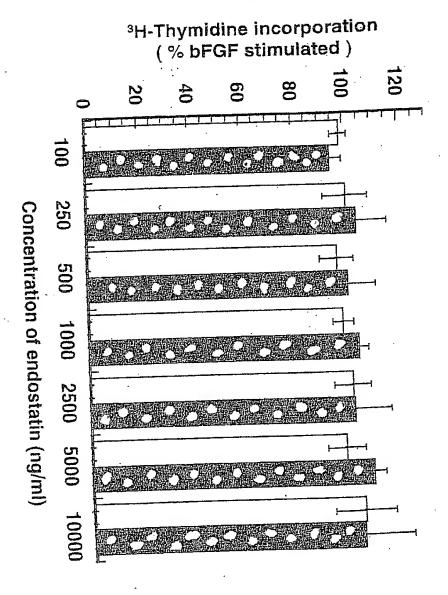
His endostatin (bacteria)
His endostatin (Yeast)
His endostatin (His endostatin (Yeast)





(†

Fig..ll



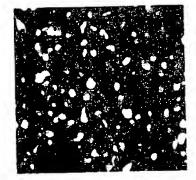
p. 31, 5 - 3

€3(₀....-

110-8291,0441

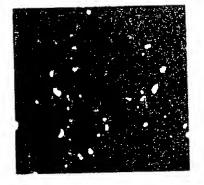
(j²)(+-

FIG. 12 A



Control +bFGF

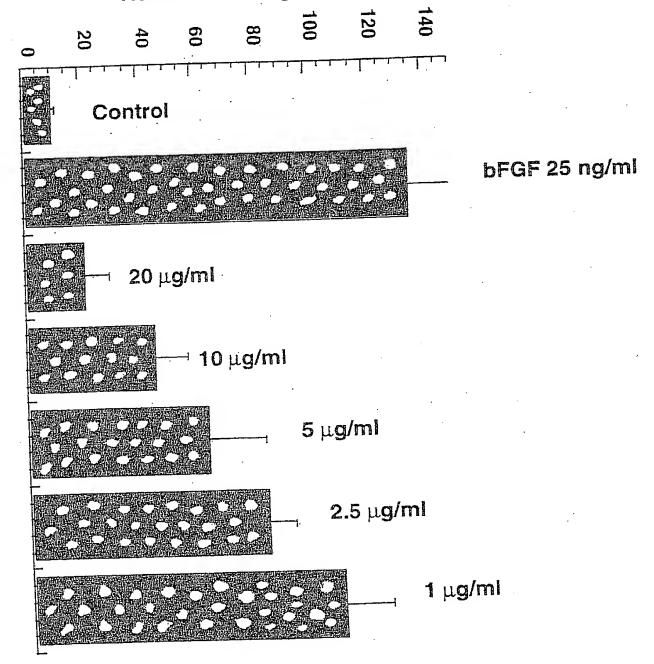
FIG.12B



Endostatin 20 μg/ml

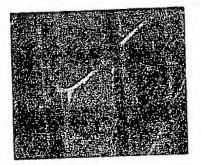
Treatment

Relative cell migration



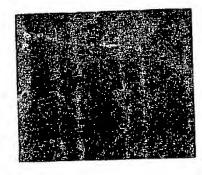
 $\left(\begin{smallmatrix} 1 & 1 & 1 \\ 1 & 2 & 1 \end{smallmatrix}\right)$

FIG.14a



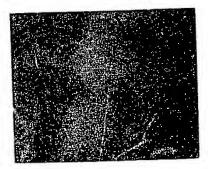
Negative control

FIG.14_b



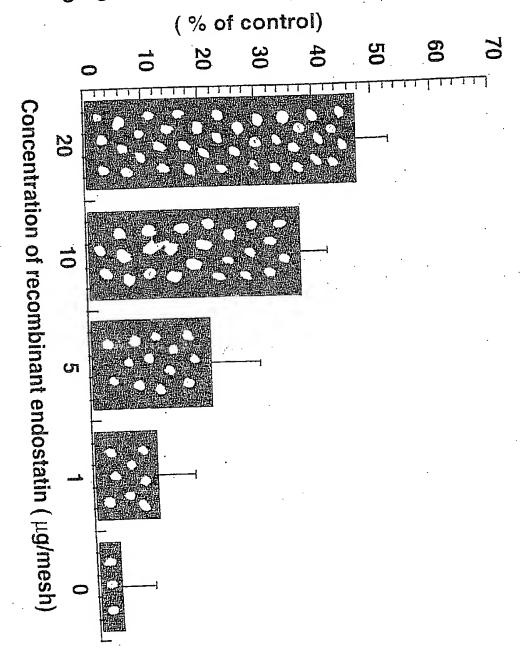
Positive control (VEGF)

FIG.14c

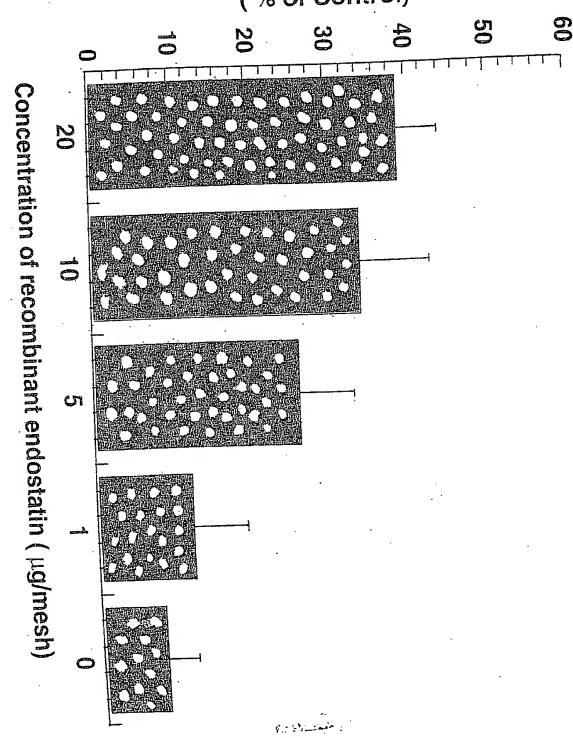


VEGF + endostatin 20 μg

Angiogenic inhibition in response to VEGF



Angiogenic inhibition in response to bFGF (% of Control)



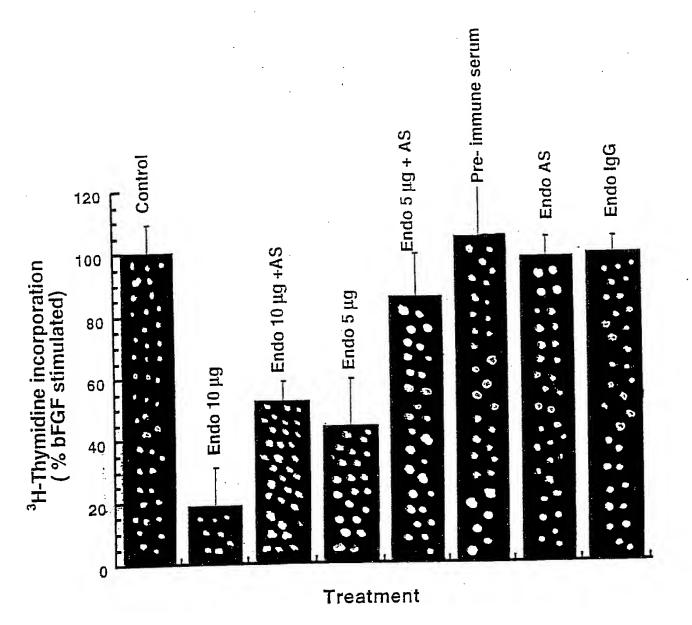
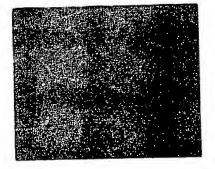


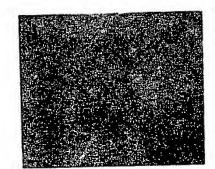
Fig. 16

FIG.17a

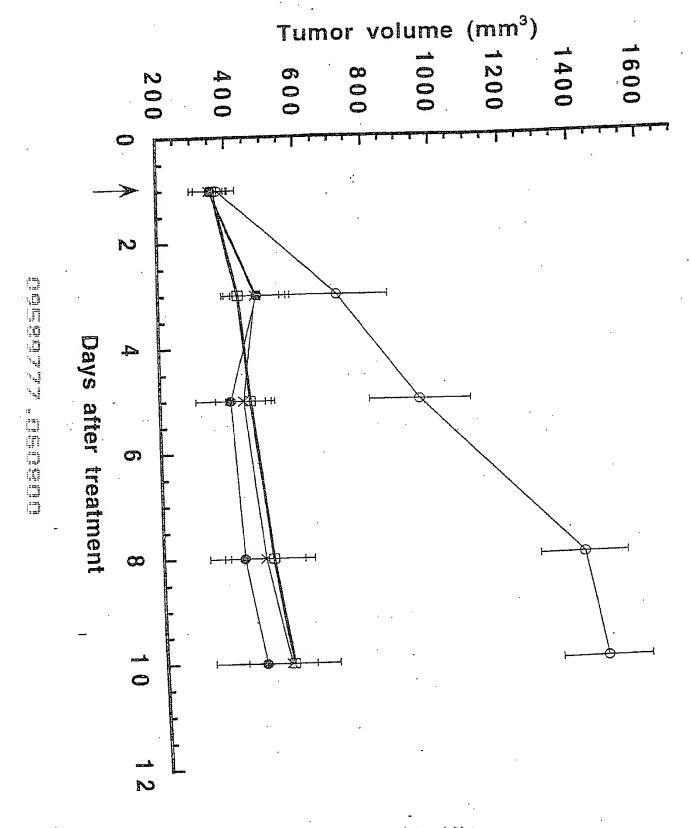


VEGF + endostatin (10 μ g)

FIG.17b



VEGF + endostatin (10 μ g) + polyclonal antiserum (50 μ g)



110-8291.84/1/

19. j.

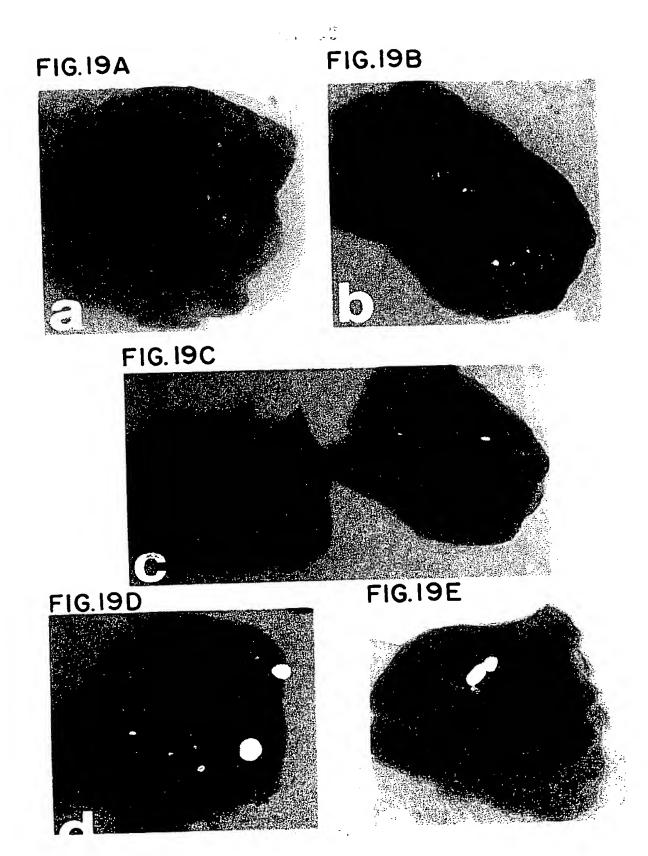
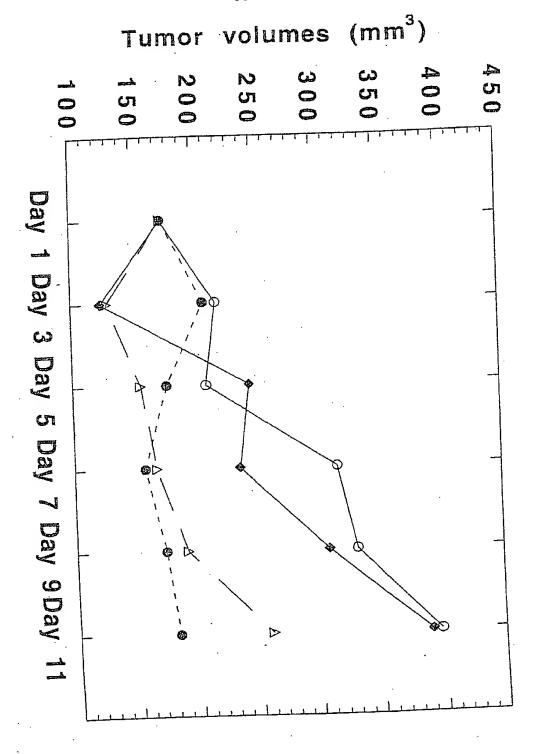
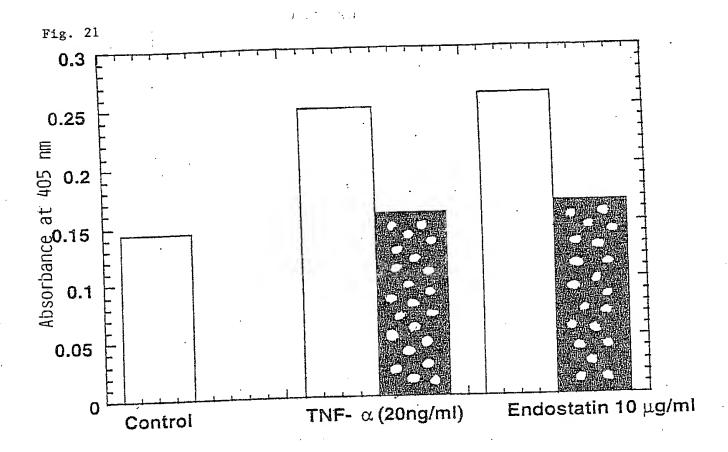
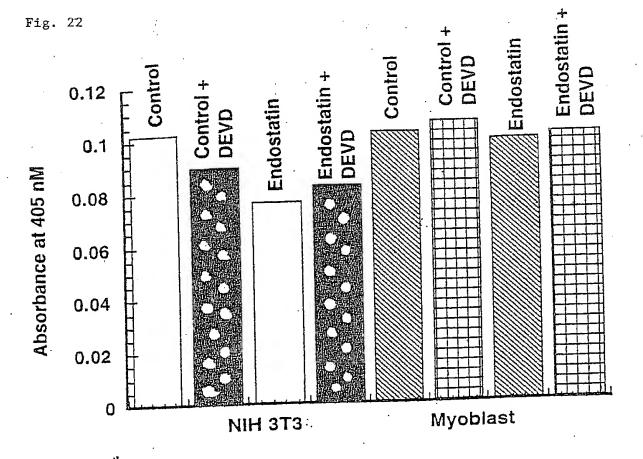


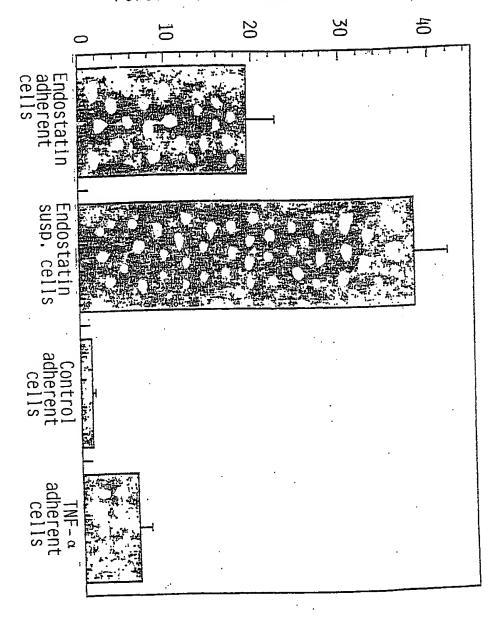
Fig. 20

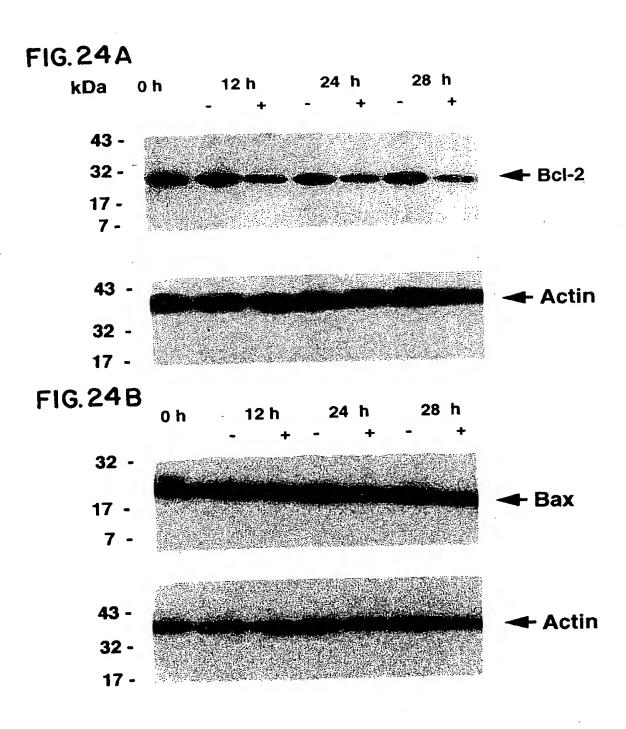


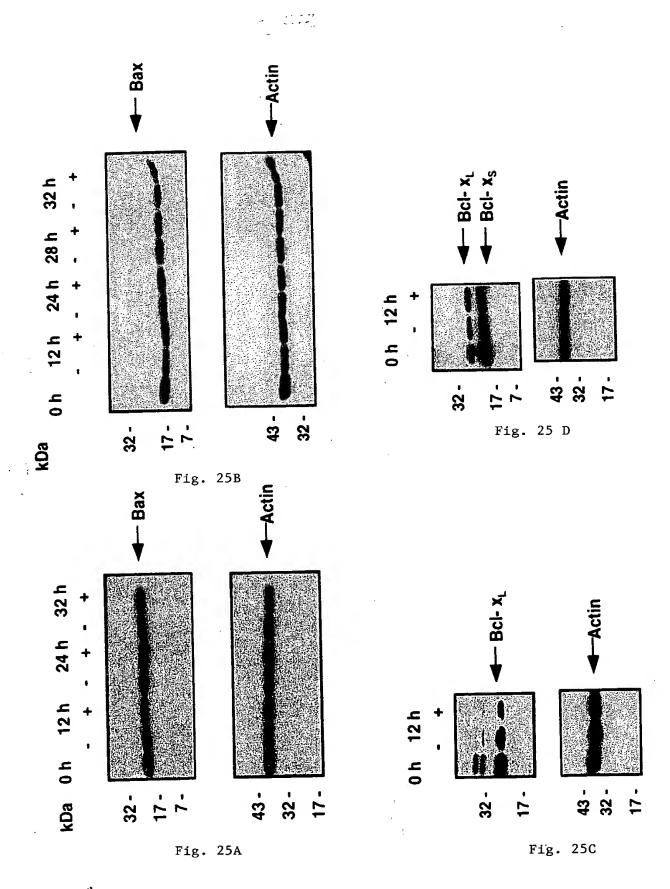




Percentage of apoptotic cells







Construct	Primer Sequence	Cloning Sites	Vector	Protein Sequence
pET17b/ his.mendo	5'-GGC ATA TGC ATA CTC ATC AGG- ACT TT-3' (up) (SEQ ID NO:3)	NdeI & XhoI	Prokaryotic expression, pET	MGHHHHHHHHHHHSSGHIDDDDKH M-mendo (SEQ ID NO:5)
	5' AAC TCG AGC TAT TTG GAG AAA- GAG GT-3' (down) (SEQ ID NO:4)			
pET28a/ mendo	5'-GGC ATA TGC ATA CTC ATC AGG- ACT TT-3' (up) (SEQ ID NO:3)	Ndel & Notl	Prokaryotic expression, pET	MGSSHHHHHHSSGLVPRGSHM- mendo (SEQ ID NO:7)
	5'-AAG CGG CCG CCT ATT TGG AGA-AAG AGG T-3' (down) (SEQ ID NO:6)			
pET28a/ EM-1	5' TTC CAT ATG CAT ACT CAT CAG- GAC TTT CAG CCA-3' (up) (SEQ ID NO:8)		Prokaryotic expression, pET	MGSSHHHHHHSSGLVPRGSHM-me ndo (SEQ ID NO:7)
	5' TTA GCG GCC GCC TAC TCA ATG- CAC AGG ACG ATG TA-3' (down) (SEQ ID NO:9)			
pET28a/ EM-2	5' TTC CAT ATG CAT ACT CAT CAG- GAC TTT CAG CCA-3' (up) (SEQ ID NO:8)		Prokaryotic expression, pET	MGSSHHHHHHSSGLVPRGSHM-me ndo (SEQ ID NO:7)
	S' TTA GCG GCC GCC TAG TTG TGG- CAG CTC GCA GCT TTC TG-3' (down) (SEQ ID NO:10)			

Construct	Primer Sequence	Cloning Sites	Vector	Protein Sequence
pPICZαA/ mendo	5' GGG AAT TCC ATA CTC ATC AGG- ACT TT-3' (up) (SEQ ID NO:11)	EcoRI & NotI	Eukaryotic expression, yeast/pPICZαA	EF-mendo
	5' AAG CGG CCG CCT ATT TGG AGA- AAG AGG T-3' (down) (SEQ ID NO:6)			
pPICZαA/ His.mendo	5' AAG AAT TCC ATC ATC ATC ATC- ATC ACA GCA GC-3' (up) (SEQ ID NO:12)	EcoRI & Notl	Eukaryotic expression, yeast/pPICZαA	EFMGHHHHHHHHHHSSGHIDDDD KHM-mendo (SEQ ID NO:13)
	5' AAG CGG CCG CCT ATT TGG AGA- AAG AGG T-3' (down) (SEQ ID NO:6)			
pPICZαA/ Hendo	S' TTT GAA TTC GCC CAC AGC CAC- CGC GAC TTC CAG CCG GTG CTC- CA-3' (up) (SEQ ID NO:14)	EcoRI & NotI	Eukaryotic expression, yeast/pPICZαA	EF-hendo
	S' AAA AGC GGC CGC CTA CTT GGA- GGC AGT CAT GAA GCT GTT CTC- AA-3' (down) (SEQ ID NO:15)			
pPICZaA/ Restin	5' TTT TTT GAA TTC ATT TCA AGT- GCC AAT TAT GAG AAG CCT GCT CTG CAT TTG-3' (up) (SEQ ID NO:16)	EcoRI & NotI	Eukaryotic (Yeast), Pichia, pPICZαA	EF-restin
·	5' AAG AAT GCG GCC GCT TAC TTC- CTA GCG TCT GTC ATG AAA CTG- TTT TCG AT-3' (down) (SEQ ID NO:17)			

Construct Name	Primer Sequence	Cloning Sites	Vector	Protein Sequence
pPICZαA/ His.Restin	5' AAT TCC ATC ACC ATC ACC ATC- ACG-3' (up) (SEQ ID NO:18)	EcoRI (oligo	Eukaryotic (Yeast), Pichia, pPICZαA	EFHHHHHHH-restin (SEQ ID NO:20)
	5' AAT TCG TGA TGG TGA TGG TGA- TGG-3' (down) (SEQ ID NO:19)	insertion)		
pET28a/ apomigren	5' TTC CAT ATG ATA TAC TCC TTT- GAT GGT CGA GAC ATA ATG ACA-3' (up) (SEQ ID NO:21)	Ndel & NotI	Prokaryotic, pET system	MGSSHHHHHHHSSGLVPRGSHM-apo migren (SEQ ID NO:7)
	5' AAT GCG GCC GCT TAC TTC CTA- GCG TCT GTC ATG AAA CTG TTT- TCG AT-3' (down) (SEQ ID NO:22)			
pPICZαA/ apomigren	S' AAG AAT TCC ATC ATC ATC ATC. ATC ACA GCA GC-3' (up) (SEQ ID NO:12)	EcoRI & NotI	Eukaryotic (Yeast), Pichia, pPICZαA	EFMGSSHHHHHHSSGLVPRGSHM-apomigren (SEQ ID NO:23)
	S' AAT GCG GCC GCT TAC TTC CTA- GCG TCT GTC ATG AAA CTG TTT- TCG AT-3' (down) (SEQ ID NO:22)	-		